

## CLAIMS

What is claimed is:

1. An isolated nucleic acid molecule selected from the group consisting of:
  - a) an isolated nucleic acid molecule which hybridizes under conditions of high stringency to a nucleic acid molecule having the complementary sequence of the nucleotide sequence of SEQ ID NO: 2, wherein said isolated nucleic acid molecule, when expressed with a molecule having the sequence of SEQ ID NO: 4 and a gene encoding an IgG1 immunoglobulin constant region, encodes a polypeptide which binds hTNF $\alpha$ ;
  - b) an isolated nucleic acid molecule which hybridizes under conditions of high stringency to a nucleic acid molecule having the complementary sequence of the nucleotide sequence of SEQ ID NO: 4, wherein said isolated nucleic acid molecule, when expressed with a molecule having the sequence of SEQ ID NO: 2 and a gene encoding an IgG1 immunoglobulin constant region, encodes a polypeptide which binds hTNF $\alpha$ ; and
  - c) a complement of an isolated nucleic acid molecule of a) or b).
  
2. An isolated nucleic acid molecule selected from the group consisting of:
  - a) an isolated nucleic acid molecule which hybridizes under conditions of high stringency to DNA having the complementary sequence of the nucleotide sequence of SEQ ID NO: 2, wherein said isolated nucleic acid molecule, when expressed with a molecule having the sequence of SEQ ID NO: 4 and a gene encoding an IgG1 immunoglobulin constant region, encodes a polypeptide which binds to hTNF $\alpha$ ; and

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reads on  
original A2  
already  
in sequence

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SEQ ID NO: 2  
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- b) an isolated nucleic acid molecule which hybridizes under conditions of high stringency to DNA having the complementary sequence of the nucleotide sequence of SEQ ID NO: 4, wherein said isolated nucleic acid molecule, when expressed with a molecule having the sequence of SEQ ID NO: 2 and a gene encoding an IgG1 immunoglobulin constant region, encodes a polypeptide which binds to hTNF $\alpha$ .
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3. An isolated nucleic acid molecule selected from the group consisting of:
- a) an isolated nucleic acid molecule which, when expressed with a molecule having the sequence of SEQ ID NO: 2 and a gene encoding an IgG1 immunoglobulin constant region, encodes a polypeptide which binds to hTNF $\alpha$ , wherein said nucleic acid molecule hybridizes under conditions of moderate stringency to a nucleic acid molecule having the complementary sequence of the nucleotide sequence of SEQ ID NO: 4;
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- b) an isolated nucleic acid molecule which, when expressed with a molecule having the sequence of SEQ ID NO: 4 and a gene encoding an IgG1 immunoglobulin constant region, encodes a polypeptide which binds to hTNF $\alpha$ , wherein said nucleic acid molecule hybridizes under conditions of moderate stringency to a nucleic acid molecule having the complementary sequence of the nucleotide sequence of SEQ ID NO: 2;
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- and
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- c) a complement of an isolated nucleic acid molecule of a) or b).
4. An isolated nucleic acid molecule selected from the group consisting of:
- a) an isolated nucleic acid molecule which, when expressed with a molecule encoding a polypeptide comprising the amino acid sequence of SEQ ID NO: 5 and a gene encoding an IgG1 immunoglobulin constant
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- region, encodes a polypeptide comprising the amino acid sequence of SEQ ID NO: 3, or a fragment thereof, which binds hTNF $\alpha$ ;
- b) an isolated nucleic acid molecule which, when expressed with a molecule encoding a polypeptide comprising the amino acid sequence of SEQ ID NO: 3 and a gene encoding an IgG1 immunoglobulin constant region, encodes a polypeptide comprising the amino acid sequence of SEQ ID NO: 5, or a fragment thereof, which binds hTNF $\alpha$ ; and
- c) a complement of the isolated nucleic acid molecule of a) or b).
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- 10 An isolated nucleic acid molecule comprising a sequence selected from the group consisting of:
- a) SEQ ID NO: 2;
- b) the complementary strand of SEQ ID NO: 2;
- c) DNA sequences that hybridize under conditions of high stringency to the complementary sequence of SEQ ID NO: 2, and which, when expressed with a molecule having the sequence of SEQ ID NO: 4 and a gene encoding an IgG1 immunoglobulin constant region, encode a polypeptide which binds hTNF $\alpha$ ; and
- 15 d) RNA sequences transcribed from the sequences of a), b), or c).
- 6.
- 20 An isolated nucleic acid molecule comprising a sequence selected from the group consisting of:
- a) SEQ ID NO: 4;
- b) the complementary strand of SEQ ID NO: 4;
- c) DNA sequences that hybridize under conditions of high stringency to the complementary sequence of SEQ ID NO: 4, and which, when expressed with a molecule having the sequence of SEQ ID NO: 2 and a gene encoding an IgG1 immunoglobulin constant region, encode a polypeptide which binds hTNF $\alpha$ ; and
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- d) RNA sequences transcribed from the sequences of a), b), or c).
7. An expression vector comprising the nucleic acid molecule according to Claim 1.
8. An expression vector comprising the nucleic acid molecule according to Claim 2.
9. ~~An expression vector comprising the nucleic acid molecule according to Claim 3.~~
10. An expression vector comprising the nucleic acid molecule according to Claim 4.
11. An expression vector comprising the nucleic acid molecule according to Claim 5.
12. An expression vector comprising the nucleic acid molecule according to Claim 6.
13. An isolated nucleic acid molecule selected from the group consisting of:
- a) an isolated nucleic acid molecule which hybridizes to a nucleic acid molecule having the complementary sequence of the nucleotide sequence of SEQ ID NO: 2 under wash conditions of wash solution of 68° C 0.1x SSC/0.1% SDS and incubation with rotation for 15 minutes at 68° C, wherein said isolated nucleic acid molecule, when expressed with a molecule having the sequence of SEQ ID NO: 4 and a gene encoding an IgG1 immunoglobulin constant region, encodes a polypeptide which binds hTNF $\alpha$ ;

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- 5           b)     an isolated nucleic acid molecule which hybridizes to a nucleic acid molecule having the complementary sequence of the nucleotide sequence of SEQ ID NO: 4 under wash conditions of wash solution of 68° C 0.1x SSC/0.1% SDS and incubation with rotation for 15 minutes at 68° C, wherein said isolated nucleic acid molecule, when expressed with a molecule having the sequence of SEQ ID NO: 2 and a gene encoding an IgG1 immunoglobulin constant region, encodes a polypeptide which binds hTNF $\alpha$ ; and
- c)     a complement of an isolated nucleic acid molecule of a) or b).
- 10   14.    An isolated nucleic acid molecule selected from the group consisting of:
- a)     an isolated nucleic acid molecule which hybridizes to DNA having the complementary sequence of the nucleotide sequence of SEQ ID NO: 2 under wash conditions of wash solution of 68° C 0.1x SSC/0.1% SDS and incubation with rotation for 15 minutes at 68° C, wherein said
- 15           isolated nucleic acid molecule, when expressed with a molecule having the sequence of SEQ ID NO: 4 and a gene encoding an IgG1 immunoglobulin constant region, encodes a polypeptide which binds hTNF $\alpha$ ;
- b)     an isolated nucleic acid molecule which hybridizes to DNA having the complementary sequence of the nucleotide sequence of SEQ ID NO: 4 under wash conditions of wash solution of 68° C 0.1x SSC/0.1% SDS, and incubation with rotation for 15 minutes at 68° C, wherein said
- 20           isolated nucleic acid molecule, when expressed with a molecule having the sequence of SEQ ID NO: 2 and a gene encoding an IgG1 immunoglobulin constant region, encodes a polypeptide which binds hTNF $\alpha$ ; and
- 25           c)     a complement of an isolated nucleic acid molecule of a) or b).

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15. An isolated nucleic acid molecule selected from the group consisting of:
- 5 a) an isolated nucleic acid molecule which, when expressed with a molecule having the sequence of SEQ ID NO: 2 and a gene encoding an IgG1 immunoglobulin constant region, encodes a polypeptide which binds hTNF $\alpha$ , wherein said nucleic acid molecule hybridizes to a nucleic acid molecule having the complementary sequence of the nucleotide sequence of SEQ ID NO: 4 under wash conditions of wash solution of 42° C 0.2x SSC/0.1% SDS and incubation with rotation for 15 minutes at 42° C;
- 10 b) an isolated nucleic acid molecule which, when expressed with a molecule having the sequence of SEQ ID NO: 4 and a gene encoding an IgG1 immunoglobulin constant region, encodes a polypeptide which binds hTNF $\alpha$ , wherein said nucleic acid molecule hybridizes to a nucleic acid molecule having the complementary sequence of the nucleotide sequence of SEQ ID NO: 2 under wash conditions of wash solution of 42° C 0.2x SSC/0.1% SDS and incubation with rotation for 15 minutes at 42° C; and
- 15 c) a complement of an isolated nucleic acid molecule of a) or b).
16. An isolated nucleic acid molecule comprising a DNA sequence that hybridizes to the complementary sequence of SEQ ID NO: 2 under wash conditions of wash solution of 68° C 0.1x SSC/0.1% SDS and incubation with rotation for 15 minutes at 68° C, said molecule, when expressed with a molecule having the sequence of SEQ ID NO: 4 and a gene encoding an IgG1 immunoglobulin constant region, encoding a polypeptide which binds hTNF $\alpha$ , or an RNA sequence transcribed from the DNA sequence.
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17. An isolated nucleic acid molecule comprising a DNA sequence that hybridizes to the complementary sequence of SEQ ID NO: 4 under wash conditions of

removes  
qualifier  
about  
binding  
TNF $\alpha$

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wash solution of 68° C 0.1x SSC/0.1% SDS and incubation with rotation for 15 minutes at 68° C, said molecule, when expressed with a molecule having the sequence of SEQ ID NO: 2 and a gene encoding an IgG1 immunoglobulin constant region, encoding a polypeptide which binds hTNF $\alpha$ , or an RNA sequence transcribed from the DNA sequence.

18. An isolated nucleic acid molecule selected from the group consisting of:
  - a) an isolated nucleic acid molecule which hybridizes under conditions of high stringency to a nucleic acid molecule having the complementary sequence of the nucleotide sequence of SEQ ID NO: 2, wherein said isolated nucleic acid molecule, when expressed with a molecule having the sequence of SEQ ID NO: 4 and a gene encoding an IgG1 immunoglobulin constant region, encodes a polypeptide which binds and inhibits hTNF $\alpha$ ;
  - b) an isolated nucleic acid molecule which hybridizes under conditions of high stringency to a nucleic acid molecule having the complementary sequence of the nucleotide sequence of SEQ ID NO: 4, wherein said isolated nucleic acid molecule, when expressed with a molecule having the sequence of SEQ ID NO: 2 and a gene encoding an IgG1 immunoglobulin constant region, encodes a polypeptide which binds and inhibits hTNF $\alpha$ ; and
  - c) a complement of an isolated nucleic acid molecule of a) or b).
19. An isolated nucleic acid molecule selected from the group consisting of:
  - a) an isolated nucleic acid molecule which hybridizes under conditions of high stringency to DNA having the complementary sequence of the nucleotide sequence of SEQ ID NO: 2, wherein said isolated nucleic acid molecule, when expressed with a molecule having the sequence of SEQ

ID NO: 4 and a gene encoding an IgG1 immunoglobulin constant region, encodes a polypeptide which binds and inhibits hTNF $\alpha$ ; and

- b) an isolated nucleic acid molecule which hybridizes under conditions of high stringency to DNA having the complementary sequence of the nucleotide sequence of SEQ ID NO: 4, wherein said isolated nucleic acid molecule, when expressed with a molecule having the sequence of SEQ ID NO: 2 and a gene encoding an IgG1 immunoglobulin constant region, encodes a polypeptide which binds and inhibits hTNF $\alpha$ .

20. An isolated nucleic acid molecule selected from the group consisting of:

- 10 a) an isolated nucleic acid molecule which, when expressed with a molecule having the sequence of SEQ ID NO: 2 and a gene encoding an IgG1 immunoglobulin constant region, encodes a polypeptide which binds and inhibits hTNF $\alpha$ , wherein said nucleic acid molecule hybridizes under conditions of moderate stringency to a nucleic acid molecule having the complementary sequence of the nucleotide sequence of SEQ ID NO: 4;
- 15 b) an isolated nucleic acid molecule which, when expressed with a molecule having the sequence of SEQ ID NO: 4 and a gene encoding an IgG1 immunoglobulin constant region, encodes a polypeptide which binds and inhibits hTNF $\alpha$ , wherein said nucleic acid molecule hybridizes under conditions of moderate stringency to a nucleic acid molecule having the complementary sequence of the nucleotide sequence of SEQ ID NO: 2; and
- 20 c) a complement of an isolated nucleic acid molecule of a) or b).

25 21. An isolated nucleic acid molecule selected from the group consisting of:

- a) an isolated nucleic acid molecule which, when expressed with a molecule encoding a polypeptide comprising the amino acid sequence of



- SEQ ID NO: 5 and a gene encoding an IgG1 immunoglobulin constant region, encodes a polypeptide comprising the amino acid sequence of SEQ ID NO: 3, or a fragment thereof, which binds and inhibits hTNF $\alpha$ ;
- 5      b)      an isolated nucleic acid molecule which, when expressed with a molecule encoding a polypeptide comprising the amino acid sequence of SEQ ID NO: 3 and a gene encoding an IgG1 immunoglobulin constant region, encodes a polypeptide comprising the amino acid sequence of SEQ ID NO: 5, or a fragment thereof, which binds and inhibits hTNF $\alpha$ ; and
- 10      c)      a complement of the isolated nucleic acid molecule of a) or b).
22.      An isolated nucleic acid molecule comprising a sequence selected from the group consisting of:
- 15      a)      SEQ ID NO: 2;
- b)      the complementary strand of SEQ ID NO: 2;
- c)      DNA sequences that hybridize under conditions of high stringency to the complementary sequence of SEQ ID NO: 2, and which, when expressed with a molecule having the sequence of SEQ ID NO: 4 and a gene encoding an IgG1 immunoglobulin constant region, encode a polypeptide which binds and inhibits hTNF $\alpha$ ; and
- 20      d)      RNA sequences transcribed from the sequences of a), b) or c).
23.      An isolated nucleic acid molecule comprising a sequence selected from the group consisting of:
- 25      a)      SEQ ID NO: 4;
- b)      the complementary strand of SEQ ID NO: 4;
- c)      DNA sequences that hybridize under conditions of high stringency to the complementary sequence of SEQ ID NO: 4, and which, when expressed with a molecule having the sequence of SEQ ID NO: 2 and a gene

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encoding an IgG1 immunoglobulin constant region, encode a polypeptide which binds and inhibits hTNF $\alpha$ ; and

- d) RNA sequences transcribed from the sequences of a), b) or c).

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